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Web Page for STN Seminar Schedule - N. America
{
m \underline{NEWS}} 2 NOV 21 CAS patent coverage to include exemplified prophetic
                 substances identified in English-, French-, German-,
                 and Japanese-language basic patents from 2004-present
NEWS 3 NOV 26 MARPAT enhanced with FSORT command
NEWS 4 NOV 26 CHEMSAFE now available on STN Easy
NEWS 5 NOV 26 Two new SET commands increase convenience of STN
                searching
NEWS 6 DEC 01 ChemPort single article sales feature unavailable
NEWS 7 DEC 12 GBFULL now offers single source for full-text
                 coverage of complete UK patent families
NEWS 8 DEC 17 Fifty-one pharmaceutical ingredients added to PS
NEWS 9 JAN 06 The retention policy for unread STNmail messages
                 will change in 2009 for STN-Columbus and STN-Tokyo
NEWS 10 JAN 07
                WPIDS, WPINDEX, and WPIX enhanced Japanese Patent
                 Classification Data
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NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3, AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

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NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS LOGIN Welcome Banner and News Items
NEWS IPC8 For general information regarding STN implementation of IPC 8
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Enter NEWS followed by the item number or name to see news on that specific topic.

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FILE 'HOME' ENTERED AT 08:58:07 ON 27 JAN 2009

=> file capluc

'CAPLUC' IS NOT A VALID FILE NAME SESSION CONTINUES IN FILE 'HOME'

Enter "HELP FILE NAMES" at an arrow prompt (=>) for a list of files that are available. If you have requested multiple files, you can specify a corrected file name or you can enter "IGNORE" to continue accessing the remaining file names entered.

=> %CA

THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE Some commands only work in certain files. For example, the EXPAND command can only be used to look at the index in a file which has an index. Enter "HELP COMMANDS" at an arrow prompt (=>) for a list of commands which can be used in this file.

=> help commands

Enter one of these commands at the arrow prompt (=>).

```
DELETE ----- Delete saved or current session items. DISPLAY ----- Display saved or current session items. FILE ------ Specify the search and display file.
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HELP ----- For help on how to use the system.

INDEX ----- Specify the Index environment.

LOGOFF ---- End the online session.

NEWS ----- Display current news about the system.

ORDER ----- Order an original document or copy.

SAVE ----- Save an L-numbered query or answer set.

SET ----- Set terminal and interaction options.

? ----- The same as HELP.

Enter one of these commands at the arrow prompt (=>).
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For more detailed information about a command, enter HELP followed by the command name at an arrow prompt.

Example: => HELP SEARCH

For information on entering multiple commands at a single arrow prompt, enter ${\tt HELP}$ COMMAND STACKING.

=> help command stacking

You can stack several commands at a single arrow prompt (=>) by separating the command steps with a semicolon. For example, a search for records containing two terms is followed by a display of the first two answers in the ALL format:

Example: => S IMMUNOASSAY AND RADIO?; D 1-2 ALL

Spaces around the semicolon are not critical. A maximum of 300 characters is permitted at a single arrow. (The maximum number of characters in a search statement is 256.)

=> 8 MCA

THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE Some commands only work in certain files. For example, the EXPAND command can only be used to look at the index in a file which has an index. Enter "HELP COMMANDS" at an arrow prompt (=>) for a list of commands which can be used in this file.

=> help search

The SEARCH command is used to execute a search in the current file.

To use this command, enter SEARCH and a query name or a logic expression. The system will create an L-number answer set containing the results of the search.

A logic expression (profile) consists of L-numbers, E-numbers, text or numeric terms and/or saved query names, joined by Boolean operators and/or by appropriate proximity operators or by numeric operators in numeric expressions.

The order of precedence for the execution of operators is (highest first): numeric operators; (W), (NOTW), (A), and (NOTA); (S) and (NOTS); (P) and (NOTP); (L) and (NOTL); AND and NOT; then OR. Parentheses (nesting) can be used to modify this order. For information on the use of operators, enter HELP OPERATORS at an arrow prompt (=>). Enter HELP NUMERIC for an explanation of how to use numeric terms in a search.

The search terms you choose must be appropriate for the file you are in, e.g., structures can be searched in the REGISTRY file but not in

the CAPLUS file. Generic structure files may be searched only with single structures, without logic operators or screen terms.

Ranges of L-numbers and/or E-numbers may be searched as if you had connected them with OR operators. For example, S E3-E6,E12,L2,L9-8 would be searched as if you had entered E3 OR E4 OR E5 OR E6 OR E12 OR L2 OR L9 OR L8.

To automatically add plurals for terms in the Basic Index or fields that comprise the Basic Index in a single search in an English language database, include PLURALS=ON in the command line, e.g., SEARCH HEDGE AND CLIPPER PLURALS=ON. For more information on searching plurals automatically, enter HELP_SET_PLURALS at an arrow prompt).

You may search a phrase in a field that contains single words and an appropriate operator, usually (W), will automatically be inserted between the words in the phrase.

Example:

=> SEARCH ACID RAIN AND POLLUTION
752118 ACID
5169 RAIN
1214 ACID RAIN
(ACID(W)RAIN)
93061 POLLUTION
L2 1214 ACID RAIN AND POLLUTION

If you do not wish to see how a phrase was actually searched, enter SET INTERPRET OFF at an arrow prompt before executing the search. For more information, enter HELP SET INTERPRET.

You may select terms from an answer set in one file and search these terms in the same or another field in the same or another file. For more information on this type of file crossover, enter HELP SMARTSELECT at an arrow prompt. For more information on other types of file crossover, enter $\underline{\text{HELP CROSSOVER}}$ in the file.

You may choose to have the SEARCH command automatically inserted into your input query. To do this, enter SET AUTOSEARCH ON at an arrow prompt. For more information, enter HELP SET AUTOSEARCH.

If a saved query appears in a search, the full name must be entered, including /Q, e.g., SEARCH L3 AND HEDGE/Q NOT SULFUR/Q.

Saved answer sets, L-number lists, and SDI profiles must have L-numbers to be used in the SEARCH command. First ACTIVATE the saved item. Then use the L-number, not the saved name, in the SEARCH command.

Searches can be done on a limited portion of the file. For an explanation, enter $\underline{\text{HELP SEARCH RANGE}}$ at an arrow prompt.

Search terms may be truncated. For information on truncation symbols, enter $\underline{\text{HELP TRUNCATION}}$ at an arrow prompt. To see what terms or symbols may need special care when used in a search, enter HELP RESERVED.

To have L-numbers assigned to intermediate postings in a SEARCH, enter

SEARCH STEPS. For more information, enter HELP SEARCH STEPS at an arrow prompt.

With fields for which a thesaurus file exists, search queries may be enriched with additional search terms such as Narrower Terms, Broader Terms, Related Terms, etc. For more information about using a thesaurus in searches, enter HELP_THESAURUS at an arrow prompt in the desired file.

When SmartSelect L-numbers are searched, a new SmartSelect L-number can be created that contains those terms for which there were no postings. For more information, enter HELP SET AUDIT at an arrow prompt.

=> help thesaurus

HELP FOR 'THESAURUS' IS NOT AVAILABLE

For information about help messages available in all files, enter "HELP MESSAGES". For information about help messages available for the current file, enter "HELP DIRECTORY". For a list of commands, enter "HELP COMMANDS".

=> help commands

Enter one of these commands at the arrow prompt (=>).

DELETE ----- Delete saved or current session items. DISPLAY ----- Display saved or current session items. FILE ----- Specify the search and display file. HELP ----- For help on how to use the system. INDEX ----- Specify the Index environment. LOGOFF ----- End the online session. NEWS ----- Display current news about the system. ORDER ----- Order an original document or copy. SAVE ----- Save an L-numbered query or answer set. SET ----- Set terminal and interaction options. ? ----- The same as HELP. Enter one of these commands at the arrow prompt (=>).

For more detailed information about a command, enter HELP followed by the command name at an arrow prompt.

Example: => HELP SEARCH

For information on entering multiple commands at a single arrow prompt, enter HELP COMMAND STACKING.

=> £il@

ENTER A FILE NAME OR (HOME): caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.66 FULL ESTIMATED COST 0.66

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FILE COVERS 1907 - 27 Jan 2009 VOL 150 ISS 5 FILE LAST UPDATED: 26 Jan 2009 (20090126/ED)

Caplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2008.

CAS Information Use Policies apply and are available at:

http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> BCV

15295 HCV 27 HCVS 1 15299 HCV

(HCV OR HCVS)

=> L1 and NS3

3283 NS3

L2 1896 L1 AND NS3

=> 12 and NS4

814 NS4

L3 280 L2 AND NS4

=> L3 and NSSb

1119 NS5B

L4 43 L3 AND NS5B

=> vector (1) 14

PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH

FIELD CODE - 'AND' OPERATOR ASSUMED 'VECTOR (L) L4'

193868 VECTOR 127643 VECTORS

265109 VECTOR

(VECTOR OR VECTORS)

L5 14 VECTOR (L) L4

=> adenovirus and L5

29681 ADENOVIRUS

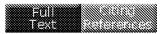
4321 ADENOVIRUSES 30502 ADENOVIRUS

(ADENOVIRUS OR ADENOVIRUSES)

L6 5 ADENOVIRUS AND L5

=> D L6 IBIB ABS 1-5

L6 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2008:1157516 CAPLUS

DOCUMENT NUMBER: 149:400310

TITLE: Compositions comprising the hepatitis C virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

antiviral vaccine uses

INVENTOR(S): Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene SA, Fr. PCT Int. Appl., 103pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

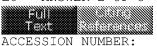
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| US 20070269460 A1 20071122 US | | | | | | | | | | US 2 | 007- | 7236 | 38 | | 2 | 0070 | 321 |
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| | | | | | | | | | | | 003- | 6772 | | | A 2 | 0030 | 605 |
| | | | | | | | | | | WO 2 | 004- | FR50: | 214 | 1 | W 2 | 0040 | 604 |
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AΒ The invention relates to the use of a therapeutically effective amt. of a peptide compn. comprising a polyprotein NS3/NS4 of the hepatitis C virus (HCV) as well as a polypeptide NS5b of the HCV, for the prepn. of a medicament for administration to a HCV-infected subject in the treatment of hepatitis C. The invention provides a compd. contg. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein NS3/NS4 and the polypeptide NS5b. Preclin. studies demonstrate that three sub-cutaneous injections of MVA vector-based NS34-NS5B construct at one week interval (weeks 1, 2 and 3) represent an optimized protocol to induce HCV specific IFNy producing T cells and cytotoxic T lymphocytes. A recall injection performed several months after the first series (either at month 4 or month 6) was able to enhance both CD4+ and CD8+ T cell responses. 8

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 2 OF 5 CAPLUS COPYRIGHT 2009 ACS on STN L6



TITLE:

2008:570045 CAPLUS

A vector-based minigene vaccine approach results in

strong induction of T-cell responses specific of

hepatitis C virus

AUTHOR(S): Martin, Perrine; Simon, Benjamin; Lone, Yu-Chun;

Chatel, Laurence; Barry, Ronald; Inchauspe, Genevieve;

Fournillier, Anne

CORPORATE SOURCE: Infectious Diseases Department, TRANSGENE SA, Lyon,

69364, Fr.

SOURCE: Vaccine (2008), 26(20), 2471-2481

CODEN: VACCDE; ISSN: 0264-410X

PUBLISHER: Elsevier Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Summary: Multiepitope-based vaccines against hepatitis C virus (HCV) were designed in the form of three minigenes encompassing four domains of the NS3, NS4 and NS5B proteins that contain multiple class I/II restricted epitopes. The polyEp-WT minigene encodes all four domains in fusion, the polyEp-C minigene encodes the same fusion but optimized for mammalian translation and the polyEp-E3 minigene has an addnl. endoplasmic reticulum targeting sequence. Whereas the minigenes vectorised by DNA were poorly immunogenic, adenovirus vectorization induced strong and broader IFNγ-ELISpot and CTL responses in HLA-A2 transgenic mice. In addn., polyEp-WT and polyEp-E3 responses were found cross-reactive in a recombinant Listeria-NS3-based surrogate challenge. This study illustrates the potency of vectorised minigenes in the field of HCV vaccine development.

REFERENCE COUNT: 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER :

ACCESSION NUMBER: 2007:1461700 CAPLUS

DOCUMENT NUMBER: 148:260241

TITLE: The Functional Evaluation of Dendritic Cell Vaccines

Based on Different Hepatitis C Virus Nonstructural

Genes

AUTHOR(S): Tian, Yuan; Zhang, Heng-Hui; Wei, Lai; Du, Shao-Cai;

Chen, Hong-Song; Fei, Ran; Liu, Feng

CORPORATE SOURCE: Hepatology Institute, Peking University People's

Hospital, Beijing, Peop. Rep. China

SOURCE: Viral Immunology (2007), 20(4), 553-561

CODEN: VIIMET; ISSN: 0882-8245

PUBLISHER: Mary Ann Liebert, Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Hepatitis C virus (HCV) nonstructural (NS) genes are relatively conserved and play crit. roles in cellular immune responses against HCV. The aim of the study was to evaluate the immunogenicity of the different HCV NS genes through transduction of DCs and presentation to T cells. Monocyte-derived DCs from healthy donors were infected with the recombinant adenovirus (Ad) harboring HCV NS3 (AdNS3), NS4 (NS4A and NS4B; AdNS4), NS5 (NS5A and NS5B; AdNS5), NS3/NS4 (AdNS3/NS4), and NS4/NS5 (AdNS4/NS5) genes, and then used to stimulate autologous lymphocytes in vitro. Antigen-specific cellular immune responses were detected by interferon-γ (IFN-γ), interleukin 4 (IL-4), and Granzyme B (GrB) enzyme-linked immunospot assays (ELISPOT). DCs expressing different HCV NS genes all induced pos. immune responses. Furthermore, DCs transfected with AdNS3/NS4 were superior to DCs infected with AdNS3 or AdNS4 in inducing HCV-specific immunity. The same results were obtained when the authors compared DCs infected with

AdNS4/NS5 to AdNS4 or AdNS5. DCs transduced with NS3/NS4 or NS4/NS5 had similar ability to elicit specific immune responses to HCV.

REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2009 ACS on STN

FOUR STORY NUMBER

ACCESSION NUMBER: 2007:1334675 CAPLUS

DOCUMENT NUMBER: 148:9402

TITLE: Compositions comprising the hepatitis C virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

vaccine uses

INVENTOR(S): Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene S.A., Fr.

SOURCE: U.S. Pat. Appl. Publ., 75 pp., Cont.-in-part of U.S.

Ser. No. 559,431.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

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| | | 2007 | 0060 | 460 | | A1 | _ | 2007 | 1100 | | | | 7026 | | | 2 | 0070 | 201 |
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| | WO | 2008 | 1136 | 06 | | A1 | | 2008 | 0925 | | WO 2 | 008- | EP23 | 00 | | 2 | 0080 | 321 |
| | | W: | ΑE, | AG, | AL, | AM, | AO, | ΑT, | AU, | AZ, | BA, | BB, | BG, | BH, | BR, | BW, | BY, | BZ, |
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| | | | ΤG, | BW, | GH, | GM , | KΕ, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, |
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AB The invention provides a compd. contg. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein NS3/NS4 and the polypeptide NS5b. The inventive compd. can be used for a therapeutic application.

L6 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2009 ACS on STN

FUI IESE

ACCESSION NUMBER: 2002:908392 CAPLUS

DOCUMENT NUMBER: 138:13314

TITLE: Comparative vaccine studies in HLA-A2.1-transgenic mice reveal a clustered organization of epitopes

presented in hepatitis C virus natural infection

AUTHOR(S): Himoudi, Nourredine; Abraham, Jean-Daniel;

Fournillier, Anne; Lone, Yu Chun; Joubert, Aurelie; Op De Beeck, Anne; Freida, Delphinc; Lemonnier, Francois;

Kieny, Marie Paule; Inchauspe, Genevieve

CORPORATE SOURCE: Unite Mixte CNRS-BioMerieux, UMR 2142, Ecole Normale

Superieure, Lyon, 69364, Fr.

SOURCE: Journal of Virology (2002), 76(24), 12735-12746

CODEN: JOVIAM; ISSN: 0022-538X

PUBLISHER: American Society for Microbiology

DOCUMENT TYPE: Journal LANGUAGE: English

AB A polyepitopic CD8+-T-cell response is thought to be crit. for control of hepatitis C virus (HCV) infection. Using transgenic mice, we analyzed the immunogenicity and dominance of most known HLA-A2.1 epitopes presented during infection by using vaccines that carry the potential to enter clin. trials: peptides, DNA, and recombinant adenoviruses. The vaccines capacity to induce specific cytotoxic T lymphocytes and interferon gamma-producing cells revealed that immunogenic epitopes are clustered in specific antigens. For two key antigens, flanking regions were shown to greatly enhance the scope of epitope recognition, whereas a DNA-adenovirus prime-boost vaccination strategy augmented epitope immunogenicity, even that of subdominant ones. The present study reveals a clustered organization of HCV immunogenic HLA.A2.1 epitopes and strategies to modulate their dominance.

REFERENCE COUNT: 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> DNA vaccine

936298 DNA

20617 DNAS

939555 DNA

(DNA OR DNAS)

74153 VACCINE

74608 VACCINES

92059 VACCINE

(VACCINE OR VACCINES)

L7 6211 DNA VACCINE

(DNA(W) VACCINE)

=> 1.7 and 1.4

L8 2 L7 AND L4

=> plasmid and L4

139820 PLASMID

53793 PLASMIDS 156869 PLASMID

(PLASMID OR PLASMIDS)

L9 9 PLASMID AND L4

=> D L8 IBIB ABS 1-2

L8 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2009 ACS on STN

TENE

ACCESSION NUMBER: 2008:1157516 CAPLUS

DOCUMENT NUMBER: 149:400310

TITLE: Compositions comprising the hepatitis C virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

antiviral vaccine uses

INVENTOR(S): Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene SA, Fr. SOURCE: PCT Int. Appl., 103pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| | PATENT I | | | KIN | D i | DATE | | | APPL | ICAT | ION 1 | NO. | | D. | ATE | | |
|-------|----------------------------------|--------------|-----|-----|-----|------|------|------|-------------|------|-------|-------|-----------------|-----|------|------|-----|
| | WO 2008 | 1136 | 06 | | A1 | _ | 2008 | 0925 | | WO 2 | 008- | EP23 | 00 | | 2 | 0080 | 321 |
| | W: | ΑE, | AG, | AL, | AM, | AO, | AΤ, | AU, | ΑZ, | BA, | BB, | BG, | BH, | BR, | BW, | BY, | BZ, |
| | | CA, | CH, | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DO, | DZ, | EC, | EE, | EG, | ES, |
| | | FI, | GB, | GD, | GE, | GH, | GM, | GΤ, | HN, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, |
| | | KG, | KM, | KN, | KΡ, | KR, | KΖ, | LA, | LC, | LK, | LR, | LS, | LT, | LU, | LY, | MA, | MD, |
| | | $	ext{ME}$, | MG, | MK, | MN, | MW, | MX, | MY, | MZ, | NA, | NG, | NΙ, | NO, | NΖ, | OM, | PG, | PH, |
| | PL, PT, | | | | RS, | RU, | SC, | SD, | SE, | SG, | SK, | SL, | SM, | SV, | SY, | ТJ, | TM, |
| | TN, TR, | | | | TZ, | UA, | UG, | US, | UZ, | VC, | VN, | ZA, | ZM, | ZW | | | |
| | RW: | AΤ, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, | FI, | FR, | GB, | GR, | HR, | HU, |
| | | IE, | IS, | ΙT, | LT, | LU, | LV, | MC, | ${ m MT}$, | ΝL, | NO, | PL, | PT, | RO, | SE, | SI, | SK, |
| | | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GQ, | GW, | \mathtt{ML} , | MR, | NE, | SN, | TD, |
| | | TG, | BW, | GH, | GM, | KE, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, |
| | | AM, | AΖ, | BY, | KG, | KΖ, | MD, | RU, | ТJ, | TM | | | | | | | |
| | <u>US 20070269460</u> A1 2007112 | | | | | | | | | US 2 | 007- | 7236. | <u> 38</u> | | 2 | 0070 | 321 |
| PRIOR | RITY APP | | | | | | US 2 | 007- | 7236 | 38 | | A2 2 | 0070 | 321 | | | |
| | | | | | | | | | | FR 2 | 003- | 6772 | | | A 2 | 0030 | 605 |
| | | | | | | | | | | WO 2 | 004- | FR50. | <u>214</u> | | W 2 | 0040 | 604 |
| | | | | | | | | | | US 2 | 005- | 5594 | <u>31</u> | | A2 2 | 0051 | 205 |
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AB The invention relates to the use of a therapeutically effective amt. of a peptide compn. comprising a polyprotein NS3/NS4 of the hepatitis C virus (HCV) as well as a polypeptide NS5b of the HCV, for the prepn. of a medicament for administration to a HCV-infected subject in the treatment of hepatitis C. The invention provides a compd. contg. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein NS3/NS4 and the polypeptide NS5b. Preclin. studies demonstrate that three sub-cutaneous injections of MVA vector-based NS34-NS5B construct at one week interval (weeks 1, 2 and 3) represent an optimized protocol to

induce HCV specific IFN γ producing T cells and cytotoxic T lymphocytes. A recall injection performed several months after the first series (either at month 4 or month 6) was able to enhance both CD4+ and CD8+ T cell responses.

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2009 ACS on STN

8

ACCESSION NUMBER: 2007:1334675 CAPLUS

DOCUMENT NUMBER: 148:9402

TITLE: Compositions comprising the hepatitis C virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

vaccine uses

INVENTOR(S): Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene S.A., Fr.

SOURCE: U.S. Pat. Appl. Publ., 75 pp., Cont.-in-part of U.S.

Ser. No. 559,431. CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

| PATENT | PATENT NO. | | | | | DATE | | | APPL | ICAT: | ION 1 | NO. | | D | ATE | | |
|----------------|------------|-----|-----|-----|-----|------|------|-------------|------|----------------|-------|-----|------|-----------------|------|-----|----|
| US 2007 | 0269 | 460 | | A1 | _ | 2007 | 1122 | | US 2 | 007- | 7236. | 38 | | 2 | 0070 | 321 | |
| FR 2855 | 758 | | | A1 | | 2004 | 1210 | | FR 2 | 003- | 6772 | | | 2 | 0030 | 605 | |
| FR 2855 | 758 | | | В1 | | 2005 | 0722 | | | | | | | | | | |
| WO 2004 | 1110 | 82 | | A2 | | 2004 | 1223 | | WO 2 | 004-1 | FR50: | 214 | | 2 | 0040 | 604 | |
| WO 2004 | 1110 | 82 | | АЗ | | 2005 | 0217 | | | | | | | | | | |
| W: | ΑE, | AG, | AL, | AM, | ΑT, | ΑU, | ΑZ, | BA, | BB, | BG, | BR, | BW, | BY, | BZ, | CA, | CH, | |
| | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | EG, | ES, | FI, | GB, | GD, | |
| | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, | KG, | KP, | KR, | KΖ, | LC, | |
| | LK, | LR, | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NA, | NΙ, | |
| | NO, | NZ, | OM, | PG, | PH, | PL, | PT, | RO, | RU, | SC, | SD, | SE, | SG, | SK, | SL, | SY, | |
| | ТJ, | TM, | TN, | TR, | TT, | TZ, | UA, | UG, | US, | UZ, | VC, | VN, | YU, | ZA, | ZM, | ZW | |
| RW: | BW, | GH, | GM, | KE, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, | AM, | |
| | ΑZ, | BY, | KG, | KΖ, | MD, | RU, | ТJ, | TM, | AT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | |
| | EE, | ES, | FI, | FR, | GB, | GR, | HU, | IE, | ΙT, | LU, | MC, | NL, | PL, | PT, | RO, | SE, | |
| | SI, | SK, | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GΑ, | GN, | GQ, | GW, | \mathtt{ML} , | MR, | NE, | |
| | SN, | TD, | TG | | | | | | | | | | | | | | |
| US 2006 | 0134 | 065 | | A1 | | 2006 | 0622 | | US 2 | 005- | 5594. | 31 | | 2 | 0051 | 205 | |
| <u>WO 2008</u> | 1136 | 06 | | A1 | | 2008 | 0925 | | WO 2 | 008 - 1 | EP23 | 00 | | 2 | 0080 | 321 | |
| W: | ΑE, | AG, | AL, | AM, | AO, | AΤ, | ΑU, | AZ, | BA, | BB, | BG, | BH, | BR, | BW, | BY, | BZ, | |
| | CA, | CH, | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DO, | DZ, | EC, | EE, | EG, | ES, | |
| | FI, | GB, | GD, | GE, | GH, | GM, | GT, | HN, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, | |
| | KG, | KM, | KN, | KP, | KR, | KΖ, | LA, | LC, | LK, | LR, | LS, | LT, | LU, | LY, | MA, | MD, | |
| | ME, | MG, | MK, | MN, | MW, | MX, | MY, | MZ, | NA, | NG, | NΙ, | NO, | NΖ, | OM, | PG, | PH, | |
| | PL, | PT, | RO, | RS, | RU, | SC, | SD, | SE, | SG, | SK, | SL, | SM, | SV, | SY, | ТJ, | TM, | |
| | TN, | TR, | TT, | TZ, | UA, | UG, | US, | UZ, | VC, | VN, | ZA, | ZM, | ZW | | | | |
| RW: | AΤ, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, | FI, | FR, | GB, | GR, | HR, | HU, | |
| | IE, | IS, | IT, | LT, | LU, | LV, | MC, | ${ m MT}$, | NL, | NO, | PL, | PT, | RO, | SE, | SI, | SK, | |
| | TR, | BF, | ΒJ, | CF, | CG, | CI, | CM, | GΑ, | GN, | GQ, | GW, | ML, | MR, | NE, | SN, | TD, | |
| | TG, | BW, | GH, | GM, | ΚE, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, | |
| | AM, | ΑZ, | BY, | KG, | KΖ, | MD, | RU, | ТJ, | TM | | | | | | | | |
| RITY APP | .: | | | | | | FR 2 | 003- | 6772 | | | A 2 | 0030 | 605 | | | |
| | | | | | | | | | | | | | | | | | |
| | | | | | | | | | WO 2 | 004: | FR50. | 214 | 1 | W 2 | 0040 | 6 | 04 |

<u>US 2005-559431</u> A2 20051205 <u>US 2007-723638</u> A2 20070321

AB The invention provides a compd. contg. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein NS3/NS4 and the polypeptide NS5b. The inventive compd. can be used for a therapeutic application.

=> D L9 IBIB ABS 1-9

L9 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

Full Text ACCESSION NUMBER:

ACCESSION NUMBER: 2008:1157516 CAPLUS

DOCUMENT NUMBER: 149:400310

TITLE: Compositions comprising the hepatitis C virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

antiviral vaccine uses

INVENTOR(S): Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene SA, Fr. SOURCE: PCT Int. Appl., 103pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| | PAT | 'ENT 1 | NO. | | | KINI | D . | DATE | | | APPL: | ICAT | ION 1 | NO. | | Di | ATE | |
|-------|-------------------------------------|--------|------|------------|-----|------|-----|------|------|-----|------------|-------|-------|------------|-------|------|------|-----|
| | WO | 2008: | 1136 | 06 | | A1 | _ | 2008 | 0925 | 1 | WO 2 | 008- | EP23 | 00 | | 2 | 0080 | 321 |
| | | W: | ΑE, | AG, | AL, | AM, | AO, | AΤ, | ΑU, | AZ, | BA, | BB, | BG, | BH, | BR, | BW, | BY, | BZ, |
| | | | CA, | CH, | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DO, | DZ, | EC, | EE, | EG, | ES, |
| | | | FI, | GB, | GD, | GE, | GH, | GM, | GT, | HN, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, |
| | | | KG, | KM, | KN, | KP, | KR, | KΖ, | LA, | LC, | LK, | LR, | LS, | LT, | LU, | LY, | MA, | MD, |
| | | | ME, | MG, | MK, | MN, | MW, | MX, | MY, | MZ, | NA, | NG, | NΙ, | NO, | NZ, | OM, | PG, | PH, |
| | | | PL, | PT, | RO, | RS, | RU, | SC, | SD, | SE, | SG, | SK, | SL, | SM, | SV, | SY, | ТJ, | TM, |
| | TN, TR, | | | | | TZ, | UA, | UG, | US, | UZ, | VC, | VN, | ZA, | ZM, | ZW | | | |
| | | RW: | AT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, | FI, | FR, | GB, | GR, | HR, | HU, |
| | | | IE, | IS, | ΙT, | LT, | LU, | LV, | MC, | MT, | NL, | NO, | PL, | PT, | RO, | SE, | SI, | SK, |
| | | | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GQ, | GW, | ML, | MR, | NE, | SN, | TD, |
| | | | TG, | BW, | GH, | GM, | KE, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, |
| | | | AM, | AZ, | BY, | KG, | KΖ, | MD, | RU, | ТJ, | TM | | | | | | | |
| | US | 20070 | 0269 | <u>460</u> | | A1 | | 2007 | 1122 | | US 2 | 007- | 7236 | <u> 38</u> | | 2 | 0070 | 321 |
| PRIO | RIORITY APPLN. INFO.: | | | | | | | | | | US 2 | 007- | 7236 | 38 | | A2 2 | 0070 | 321 |
| | | | | | | | | | | | FR 2 | 003- | 6772 | | | A 2 | 0030 | 605 |
| | | | | | | | | | | 1 | WO 2 | 004- | FR50: | 214 | 1 | W 2 | 0040 | 604 |
| | | | | | | | | | | | US 2 | 005- | 5594. | 31 | | A2 2 | 0051 | 205 |
| 7\ T) | The invention melates to the use of | | | | | | | | | | h o 20 o 1 | 0011+ | 1 001 | 1 | ffoo. | + | - m+ | o f |

AB The invention relates to the use of a therapeutically effective amt. of a peptide compn. comprising a polyprotein NS3/NS4 of the hepatitis C virus (HCV) as well as a polypeptide NS5b of the HCV, for the prepn. of a medicament for administration to a HCV-infected subject in the treatment of hepatitis C. The invention provides a compd. contg. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression

vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein ${\tt NS3/NS4}$ and the polypeptide ${\tt NS5b}$. Preclin. studies demonstrate that three sub-cutaneous injections of MVA vector-based NS34-NS5B construct at one week interval (weeks 1, 2 and 3) represent an optimized protocol to induce ${\tt HCV}$ specific IFN γ producing T cells and cytotoxic T lymphocytes. A recall injection performed several months after the first series (either at month 4 or month 6) was able to enhance both CD4+ and CD8+ T cell responses.

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 2 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

8

FUL Text

ACCESSION NUMBER:

DOCUMENT NUMBER: 148:9402

TITLE:

Compositions comprising the hepatitis ${\tt C}$ virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

vaccine uses

INVENTOR(S):

Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene S.A., Fr.

SOURCE:

U.S. Pat. Appl. Publ., 75 pp., Cont.-in-part of U.S.

Ser. No. 559,431.

2007:1334675 CAPLUS

CODEN: USXXCO

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 3

| PATENT : | NO. | | | KIN | D | DATE | | | APPL | ICAT | ION 1 | NO. | | Di | ATE | |
|-------------------------------|------|------------|-------|----------------|-----|--------------------------|------|-----|--------------|------|-------|-----|-----|-----|---------------|-----|
| US 2007 FR 2855 FR 2855 | 758 | 460 460 | | A1 A1 B1 | | 2007 2004 2005 | 1210 | | US 2 FR 2 | | | | | _ | 0070: 0030 | |
| WO 2004 WO 2004 | 1110 | | | A2 A3 | | 2005 2004 2005 | 1223 | | WO 2 | 004- | FR50. | 214 | | 2 | 0040 | 604 |
| W: | • | • | • | • | • | • | AZ, | • | • | • | • | • | • | • | • | • |
| | | | | | • | | DK, | | | • | | | | | | |
| | • | , | , | • | • | • | IL, | • | • | , | , | • | , | • | • | , |
| | • | • | • | • | • | • | MA, | • | • | • | • | • | | • | • | • |
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| DM. | , | TM, | • | - | - | - | UA, | - | - | | • | - | | - | - | |
| KW: | • | • | • | • | • | • | MZ, | • | • | • | • | • | • | • | • | • |
| | • | | | | • | • | TJ, | | • | • | | • | | • | • | • |
| | • | | | | | • | HU, | | • | | | • | • | | | • |
| | | TD, | | Dr, | ъо, | Cr, | CG, | CΙ, | CM, | GA, | GIV, | GQ, | GW, | М., | MK, | NE, |
| US 2006 | , | • | | A 1 | | 2006 | 0622 | | ric o | 005 | 5507 | 21 | | 2 | 0051: | 205 |
| WO 2008 | | | | A1 | | 2008 | | | WO 2 | | | | | | 0080: | |
| | | | Z\ T. | | | | AU, | | | | | | BB | | | |
| ٠, ٧٧٠ | • | • | • | • | • | • | CZ, | • | • | • | • | • | • | • | • | • |
| | • | • | • | • | • | • | GT, | • | • | | • | • | • | • | • | • |
| | • | • | • | • | • | • | LA, | • | • | • | • | • | • | • | • | • |
| | • | • | • | • | • | • | MY, | • | • | • | • | • | • | • | • | • |
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| RW: | • | • | • | • | • | • | DE, | • | • | • | • | • | | GR. | HR. | HU, |
| | • | • | • | • | • | • | MC, | • | • | | • | • | | • | • | |
| | • | • | • | • | • | • | CM, | • | • | • | • | • | • | • | • | • |

AB The invention provides a compd. contg. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein NS3/NS4 and the polypeptide NS5b. The inventive compd. can be used for a therapeutic application.

L9 ANSWER 3 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

FUIL TEXT

ACCESSION NUMBER: 2006:333454 CAPLUS

DOCUMENT NUMBER: 144:357638

TITLE: Application of a transgenic mouse model of hepatitis c

virus (HCV) infection and identification of

antiviral agent for ${f HCV}$ therapeutics

INVENTOR(S): Sallberg, Matti; Frelin, Lars

PATENT ASSIGNEE(S): Tripep AB, Swed.

SOURCE: PCT Int. Appl., 165 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|--|--|---|--|
| WO 2006021896 WO 2006021896 | A2 A3 | 20060302 20060817 | WO 2005-IB3736 | 20050826 |
| W: AE, AG, CN, CO, GE, GH, LC, LK, NG, NI, | AL, AM, AT, CR, CU, CZ, GM, HR, HU, LR, LS, LT, NO, NZ, OM, SY, TJ, TM | AU, AZ, DE, DK, ID, IL, LU, LV, PG, PH, | BA, BB, BG, BR, BW, DM, DZ, EC, EE, EG, IN, IS, JP, KE, KG, MA, MD, MG, MK, MN, PL, PT, RO, RU, SC, TT, TZ, UA, UG, US, | ES, FI, GB, GD, KM, KP, KR, KZ, MW, MX, MZ, NA, SD, SE, SG, SK, |
| IS, IT, CF, CG, GM, KE, | LT, LU, LV, CI, CM, GA | MC, NL, GN, GQ, NA, SD, | DK, EE, ES, FI, FR, PL, PT, RO, SE, SI, GW, ML, MR, NE, SN, SL, SZ, TZ, UG, ZM, | SK, TR, BF, BJ, TD, TG, BW, GH, |
| EP 1781690 | A2 | 20070509 | EP 2005-810181 | 20050826 |
| · | | | DK, EE, ES, FI, FR, NL, PL, PT, RO, SE, | |
| WO 2006109196 | A2 | 20061019 | WO 2006-IB1668 | 20060203 |
| WO 2006109196 | A3 | 20070315 | | |
| CN, CO, GE, GH, KZ, LC, MZ, NA, | CR, CU, CZ GM, HR, HU LK, LR, LS NG, NI, NO | DE, DK, , ID, IL, , LT, LU, , NZ, OM, | BA, BB, BG, BR, BW, DM, DZ, EC, EE, EG, IN, IS, JP, KE, KG, LV, LY, MA, MD, MG, PG, PH, PL, PT, RO, TN, TR, TT, TZ, UA, | ES, FI, GB, GD, KM, KN, KP, KR, MK, MN, MW, MX, RU, SC, SD, SE, |
| , , | ZA, ZM, ZW BG, CH, CY | , CZ, DE, | DK, EE, ES, FI, FR, | GB, GR, HU, IE, |

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IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ,
            CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
            GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM
    US 20080295185
                        Α1
                               20081127
                                           US 2008-660878
                                                                  20080506
PRIORITY APPLN. INFO.:
                                           US 2004-605030P
                                                               P 20040827
                                           US 2005-649975P
                                                               P 20050204
                                           WO 2005-IB3736
                                                              W 20050826
                                           US 2005-740362P
                                                              P 20051128
```

AB Disclosed herein is the discovery of novel NS3/4A compns. with enhanced expression abilities. Embodiments of the invention include codon optimized NS3/4A compns. and compns. with the Semliki forest virus replicon. Addnl. embodiments include transgenic organisms contg. these NS3/4A compns., methods of using these transgenic mice to screen and refine drugs, and the drugs refined by these methods. Addnl. embodiments include protease activity dependent mols. that can indicate the presence or absence of a protease inhibitor.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 4 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

3

FUI Text

ACCESSION NUMBER: 2005:303181 CAPLUS

DOCUMENT NUMBER: 142:372468

TITLE: HCV fusion proteins with modified NS3 domains and

uses thereof as immunogens

INVENTOR(S): Houghton, Michael

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S.

Ser. No. 721,479. CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|----|----------|
| | | | | - | |
| US 20050074465 | A1 | 20050407 | US 2003-612884 | | 20030702 |
| <u>US 6986892</u> | B1 | 20060117 | US 2000-721479 | | 20001122 |
| US 20060057164 | A1 | 20060316 | US 2005-195009 | | 20050802 |
| US 7449566 | В2 | 20081111 | | | |
| JP 2006265267 | A | 20061005 | JP 2006-174595 | | 20060623 |
| PRIORITY APPLN. INFO.: | | | US 1999-167502P | P | 19991124 |
| | | | US 2000-721479 | Α2 | 20001122 |
| | | | US 2002-393694P | P | 20020702 |
| | | | US 2002-394510P | P | 20020708 |
| | | | JP 2004-519849 | АЗ | 20030702 |

The disclosed invention provides hepatitis C virus (HCV) fusion proteins that include a mutated NS3 protease domain, fused to at least one other HCV epitope derived from another region of the HCV polyprotein. The fusions can be used in stimulation of a cellular immune response to HCV, such as activating hepatitis C virus (HCV)-specific T cells, including CD4+ and CD8+ T cells. The method can be used in model systems to develop HCV-specific immunogenic compns., as well as to immunize a mammal against HCV. In expts. with Rhesus macaques, the immunization with plasmid DNA encoding an NS3 (modified) NS4NS5aCore fusion protein led to activation of HCV-specific CD8-pos. T cells expressing interferon y and proliferation of HCV-specific CD4-pos. T cells. Also

presented is the use of alphavirus replicon particles.

L9 ANSWER 5 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

Texts

ACCESSION NUMBER:

ACCESSION NUMBER: 2004:392569 CAPLUS

DOCUMENT NUMBER: 140:390291

TITLE: Activation of **HCV**-specific T cells using fusion

protein vaccines comprising HCV NS3, NS4, NS5a,

and **NS5b** polypeptides

INVENTOR(S): Houghton, Michael; Coates, Steve; Selby, Mark;

Paliard, Xavier

PATENT ASSIGNEE(S): Chiron Corporation, USA SOURCE: PCT Int. Appl., 136 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PAT | PATENT NO. | | | | | D | DATE | | | APPL | ICAT | ION 1 | NO. | | D. | ATE | |
|---------------------|------------|------|------|-----|------|------|------|------|-----|------|------|-------|------------|-----|-----|------|-----|
| | 2004 | | | | | | | | | WO 2 | 003- | US33 | <u>610</u> | | 2 | 0031 | 024 |
| | W: | AE, | AG, | AL, | AM, | AT, | AU, | AZ, | BA, | BB, | BG, | BR, | BY, | BZ, | CA, | CH, | CN, |
| | | | | | | | DK, | | | | | | | | | | |
| | | | | | | | IL, | | | | | | | | | | |
| | | | | • | | • | MA, | • | | | • | • | | | • | • | • |
| | | OM, | PG, | PH, | PL, | PT, | RO, | RU, | SC, | SD, | SE, | SG, | SK, | SL, | SY, | ТJ, | TM, |
| | | TN, | TR, | TT, | TZ, | UA, | UG, | US, | UZ, | VC, | VN, | YU, | ZA, | ZM, | ZW | | |
| | RW: | GH, | GM, | KE, | LS, | MW, | MZ, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, | AM, | AZ, | BY, |
| | | KG, | KΖ, | MD, | RU, | ТJ, | TM, | ΑT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, |
| | | FI, | FR, | GB, | GR, | HU, | IE, | ΙT, | LU, | MC, | NL, | PT, | RO, | SE, | SI, | SK, | TR, |
| | | BF, | ВJ, | CF, | CG, | CI, | CM, | GΑ, | GN, | GQ, | GW, | ML, | MR, | NE, | SN, | TD, | TG, |
| | | AP, | EA, | EP, | OA | | | | | | | | | | | | |
| CA | 2505 | 611 | | | A1 | | 2004 | 0513 | | CA 2 | 003- | 2505 | 611 | | 2 | 0031 | 024 |
| AU | 2003 | 2871 | 88 | | A1 | | 2004 | 0525 | | AU 2 | 003- | 2871 | 88 | | 2 | 0031 | 024 |
| EΡ | | 2005 | 0921 | | EP 2 | 003- | 7813 | 68 | | 2 | 0031 | 024 | | | | | |
| | R: | ΑT, | BE, | CH, | DE, | DK, | ES, | FR, | GB, | GR, | ΙT, | LI, | LU, | NL, | SE, | MC, | PT, |
| | | IE, | SI, | LT, | LV, | FI, | RO, | MK, | CY, | AL, | TR, | BG, | CZ, | EE, | HU, | SK | |
| ORITY APPLN. INFO.: | | | | | | | | | | US 2 | 002- | 2813 | 41 | Ź | A 2 | 0021 | 025 |
| | | | | | | | | | | WO 2 | 003- | US33 | <u>610</u> | Ţ | W 2 | 0031 | 024 |

AB The invention provides a method of activating hepatitis C virus (HCV)-specific T cells, including CD4+ and CD8+ T cells. HCV-specific T cells are activated using fusion protein vaccines comprising HCV NS3, NS4, NS5a, and NS5b polypeptides, polynucleotides encoding such fusion proteins, or polypeptide or polynucleotide compns. contg. the individual components of these fusions. The method can be used in model systems to develop HCV-specific immunogenic compns., as well as to immunize a mammal against HCV.

L9 ANSWER 6 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

Full Text

ACCESSION NUMBER: 2002:908392 CAPLUS

DOCUMENT NUMBER: 138:13314

TITLE: Comparative vaccine studies in HLA-A2.1-transgenic mice reveal a clustered organization of epitopes

presented in hepatitis C virus natural infection

AUTHOR(S): Himoudi, Nourredine; Abraham, Jean-Daniel;

Fournillier, Anne; Lone, Yu Chun; Joubert, Aurelie; Op De Beeck, Anne; Freida, Delphinc; Lemonnier, Francois;

Kieny, Marie Paule; Inchauspe, Genevieve

CORPORATE SOURCE: Unite Mixte CNRS-BioMerieux, UMR 2142, Ecole Normale

Superieure, Lyon, 69364, Fr.

SOURCE: Journal of Virology (2002), 76(24), 12735-12746

CODEN: JOVIAM; ISSN: 0022-538X

PUBLISHER: American Society for Microbiology

DOCUMENT TYPE: Journal LANGUAGE: English

AB A polyepitopic CD8+-T-cell response is thought to be crit. for control of hepatitis C virus (HCV) infection. Using transgenic mice, we analyzed the immunogenicity and dominance of most known HLA-A2.1 epitopes presented during infection by using vaccines that carry the potential to enter clin. trials: peptides, DNA, and recombinant adenoviruses. The vaccines capacity to induce specific cytotoxic T lymphocytes and interferon gamma-producing cells revealed that immunogenic epitopes are clustered in specific antigens. For two key antigens, flanking regions were shown to greatly enhance the scope of epitope recognition, whereas a DNA-adenovirus prime-boost vaccination strategy augmented epitope immunogenicity, even that of subdominant ones. The present study reveals a clustered organization of HCV immunogenic HLA.A2.1 epitopes and strategies to modulate their dominance.

REFERENCE COUNT: 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 7 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

FUII
TEXT

ACCESSION NUMBER:

INVENTOR(S):

ACCESSION NUMBER: 2002:716438 CAPLUS

DOCUMENT NUMBER: 137:227663

TITLE: Hepatitis C virus (HCV) cDNA-based hepatocyte cell

culture system for synthesis of infectious HCV, and

uses for antiviral screening Dasgupta, Asim; Koka, Prasad S.

PATENT ASSIGNEE(S): The Regents of the University of California, USA

SOURCE: PCT Int. Appl., 42 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

| PATENT | NO. | | | KIN | D | DATE | | | APPL | ICAT | ION 1 | NO. | | Di | ATE | |
|----------------|-------------|-----------|-----|-----------------|------------------------------------|------|------|------|------|------|-------|-----|------|-----|------|-----|
| WO 2002 | | | | A2 | | 2002 | | : | WO 2 | 002- | US75 | 16 | | 2 | 0020 | 311 |
| WO 2002 | <u>0727</u> | <u>76</u> | | A3 | | 2004 | 0205 | | | | | | | | | |
| W: | ΑE, | AG, | AL, | AM, | ΑT, | ΑU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | ΒZ, | CA, | CH, | CN, |
| | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | ES, | FI, | GB, | GD, | GE, | GH, |
| | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | ΚE, | KG, | KΡ, | KR, | KΖ, | LC, | LK, | LR, |
| | LS, | LT, | LU, | LV, | 7, MA, MD, MG, 1 J, SD, SE, SG, | | | MK, | MN, | MW, | MX, | MZ, | NO, | NZ, | OM, | PH, |
| | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TN, | TR, | TT, | TZ, |
| | UA, | UG, | US, | UZ, | VN, | YU, | ZA, | ZM, | ZW | | | | | | | |
| RW: | GH, | GM, | ΚE, | LS, | MW, | MΖ, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, | AM, | AZ, | BY, |
| | KG, | KΖ, | MD, | RU, | ТJ, | TM, | ΑT, | BE, | CH, | CY, | DE, | DK, | ES, | FI, | FR, | GB, |
| | GR, | IE, | ΙT, | LU, | MC, | NL, | PT, | SE, | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GΑ, |
| | GN, | GQ, | GW, | \mathtt{ML} , | LU, MC, NL, PT, ML, MR, NE, SN, | | | | ΤG | | | | | | | |
| CA 2440433 | | | | Al | | 2002 | 0919 | | CA 2 | 002- | 2440 | 433 | | 2 | 0020 | 311 |
| <u>AU 2002</u> | | A1 | | 2002 | 0924 | | AU 2 | 002- | 2541 | 90 | | 2 | 0020 | 311 | | |
| US 20020197277 | | | | A1 | | 2002 | 1226 | | US 2 | 002- | 9603 | 9 | | 2 | 0020 | 311 |

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B2 20070227
A2 20040526 <u>EP 2002-723409</u>
     US 7183095
     EP 1421222
                                                                       20020311
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     JP 2004537279 T
                              20041216 <u>JP 2002-571</u>832
                                                                       20020311
     CN 1592794
                           Α
                                 20050309
                                              CN 2002-806237
                                                                       20020311
                                              <u>US 2001-274709P</u> P 20010309

<u>WO 2002-US7516</u> W 20020311
PRIORITY APPLN. INFO.:
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The present invention presents a method of synthesizing infectious AB hepatitis C virus (HCV) by transfecting hepatocyte cells with a gene encoding HCV and then exposing uninfected cells to the HCV to form addnl. HCV. The invention relates to a HCV cDNA-based culture system capable of synthesis of infectious HCV in cell culture and cell-to-cell spread of the virus. The expression of T7 RNA polymerase in the cytoplasm was used to transcribe the HCV cDNA under the T7 promoter to generate high quantities of HCV RNA. The viral RNA proved to be translated to produce viral structural (core, El, E2 and p7) and nonstructural (NS2, NS3, NS4A and B, NS5A and B) proteins. Viral RNA replication directed by the RNA-dependent RNA polymerase (NS5B) would then occur. Progeny virions were made and secreted into the tissue culture media, and infection of neighboring cells resulting in cell-to-cell spread of virus was demonstrated. The invention also relates to a method of measuring the level of HCV infection in a hepatocyte cell. A method for identifying a modulator of HCV activity is also presented, and a method for modulating HCV activity. The invention provides a reliable system for both genetic anal. of the viral genome and for the development of novel antiviral strategies.

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 8 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

EU Ieae

ACCESSION NUMBER: 2001:319922 CAPLUS

DOCUMENT NUMBER: 134:325205

TITLE: Activation of HCV-specific T cells using hepatitis C

virus nonstructural proteins, either alone or as

fusions

INVENTOR(S): Paliard, Xavier; Houghton, Michael; Selby, Mark

PATENT ASSIGNEE(S): Chiron Corp., USA

SOURCE: PCT Int. Appl., 56 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

| PATENT | NO. | | | KIN | D i | DATE | | | APPL | ICAT | ION 1 | NO. | | Di | ATE | |
|-------------------------------|------------|-----|---|----------------|------|----------------------|------|-----|------|------------|-------|-------|-------|----|------------|---------|
| WO 2001 WO 2001 WO 2001 | 0308 | 12 | | A2 A3 A9 | | 2001 2002 2002 | 0228 | 1 | wo 2 | 000- | US29 | 594 | | 21 | 0001 | 027 |
| | AE, | AG, | • | AM, | AT, | AU, | AZ, | • | • | • | • | • | • | • | • | • |
| | • | • | • | • | • | • | • | • | • | • | • | • | • | • | GM, LS, | • |
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| DM• | YU, GH, | ZA, | | T.S | Mīaī | M <i>7</i> | GD. | QT. | 97 | Ф <i>7</i> | ПС | 7 TaT | 7\ TT | BE | СП | CV |
| 1/1/1 | • | • | • | • | • | • | • | • | • | • | • | • | • | • | BF, | • |

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CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    CA 2389206 A1 20010503 CA 2000-2389206
                                                           20001027
    EP 1232267
                      A2
                           20020821
                                      EP 2000-973922
       R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
           IE, SI, LT, LV, FI, RO, MK, CY, AL
                          20030408 JP 2001-533809
    JP 2003512826 T
                                                           20001027
                     B1 20030513
A1 20030911
                                      US 2000-698874
    US 6562346
                                                           20001027
    US 20030170274
                                     US 2003-357619
                                                           20030203
    US 7285539
                     B2 20071023
    US 20040057960
                     A1 20040325
                                      US 2003-643679
    US 20040191767 A1 20040930
                                                       20040412
P 19991027
                                      US 2004-822607
PRIORITY APPLN. INFO.:
                                       US 1999-161713P
                                       US 2000-698874
                                       WO 2000-US29594
                                                        W 20001027
                                                      A3 20030203
                                       US 2003-357619
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AΒ The invention provides a method of activating hepatitis C virus (HCV) specific T cells, including CD4+ and CD8+ T cells. HCV-specific T cells are activated using fusion proteins comprising HCV NS3, NS4, NS5a, and NS5b polypeptides, polynucleotides encoding such fusion proteins, or polypeptide or polynucleotide compns. contg. the individual components of these fusions. The method can be used in model systems to develop HCV-specific immunogenic compns., as well as to immunize a mammal against **HCV**.

REFERENCE COUNT: THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 9 OF 9 CAPLUS COPYRIGHT 2009 ACS on STN

Text ACCESSION NUMBER:

1999:113845 CAPLUS

DOCUMENT NUMBER: 130:163166

TITLE: Test vectors containing hepatitis C or human

cytomegalovirus nucleic acid and indicator gene and methods for determining antiviral susceptibility and

resistance and for antiviral screening

Capon, Daniel J.; Whitcomb, Jeannette M.; Parkin, Neil INVENTOR(S):

Τ.

Virologic, Inc., USA PATENT ASSIGNEE(S): SOURCE: PCT Int. Appl., 128 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

| PA' | TENT : | NO. | | | KIN | D | DATE | | | APPL | ICAT | ION 1 | NO. | | D. | ATE | |
|-----|--------|---------|-----|-----|-----|-------------|----------|------|-----|----------|------|----------|----------|-----|-----|-------------------|-----|
| WO. | 9906 | 597 | | | A1 | _ | 1999 | 0211 | 1 | WO 1 | 998- | US15 | 967 | | 1 | 9980 [.] | 730 |
| | W: | AL, | AM, | AΤ, | AU, | AZ, | BA, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CU, | CZ, | DE, |
| | | DK, | EE, | ES, | FI, | GB, | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IS, | JP, | KΕ, | KG, |
| | | KP, | KR, | KΖ, | LC, | LK, | LR, | LS, | LT, | LU, | LV, | MD, | MG, | MK, | MN, | MW, | MX, |
| | | NO, | NZ, | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, |
| | | UA, | UG, | UZ, | VN, | YU, | ZW | | | | | | | | | | |
| | RW: | GH, | GM, | ΚE, | LS, | MW, | SD, | SZ, | UG, | ZW, | AT, | BE, | CH, | CY, | DE, | DK, | ES, |
| | | FI, | FR, | GB, | GR, | IE, | ΙT, | LU, | MC, | NL, | PT, | SE, | BF, | ВJ, | CF, | CG, | CI, |
| | | CM, | GΑ, | GN, | GW, | ${ m ML}$, | MR, | NE, | SN, | TD, | TG | | | | | | |
| CA | 2298 | 102 | | | Al | | 1999 | 0211 | | CA 1 | 998- | 2298 | 102 | | 1 | 9980 | 730 |
| AU | 9888 | 976 | | | Α | | 1999 | 0222 | | AU 1 | 998- | 8897 | <u>6</u> | | 1 | 9980 | 730 |
| EΡ | 1012 | 334 | | | A1 | | 2000 | 0628 | | EP 1 | 998- | 9407 | 79 | | 1 | 9980' | 730 |
| | R: | AT. | BE. | CH. | DE. | DK. | ES. | FR. | GB. | GR. | IT. | LI. | LU. | NL. | SE. | MC. | PT. |

IE, FI

JP 2001512036

PRIORITY APPLN. INFO.:

T 20010821 <u>JP 2000-505336</u> US 1997-903507

19980730 A 19970730 W 19980730

MO 1998-US15967 W 19980730

AB This invention provides a method for detg. susceptibility for an HCV or HCMV anti-viral drug comprising: (a) introducing a resistance test vector comprising a patient-derived segment and an indicator gene into a host cell; (b) culturing the host cell from (a); (c) measuring expression of the indicator gene in a target host cell, and (d) comparing the expression of the indicator gene from (c) with the expression of the indicator gene measured when steps (a-c) are carried out in the absence of the anti-viral drug, wherein a test concn. of the anti-viral drug is present at steps (a-c); at steps (b-c); or at step (c). This invention also provides a method for detg. HCV or HCMV anti-viral drug resistance in a patient

comprising: (a) detg. anti-viral drug susceptibility in the patient at a first time using the susceptibility test described above, wherein the patient-derived segment is obtained from the patient at about said time; (b) detg. anti-viral drug susceptibility of the same patient at a later time; and (c) comparing the anti-viral drug susceptibilities detd. in step (a) and (b), wherein a decrease in anti-viral drug susceptibility at the later time compared to the first time indicates development or progression of anti-viral drug resistance in the patient. This invention also provides a method for evaluating the biol. effectiveness of a candidate HCV or HCMV anti-viral drug compd. Compns. including resistance test vectors comprising a patient-derived segment comprising an HCV or HCMV gene and an indicator gene and host cells transformed with the resistance test vectors are provided.

REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> D L5 IBIB ABS 1-14

L5 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

EUI IEXE

ACCESSION NUMBER: 2008:1157516 CAPLUS

DOCUMENT NUMBER: 149:400310

TITLE: Compositions comprising the hepatitis C virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

antiviral vaccine uses

INVENTOR(S): Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene SA, Fr. SOURCE: PCT Int. Appl., 103pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

| PATENT NO. | | | | | KIN | D i | DATE | | APPLICATION NO. | | | | | | DATE | | | |
|---------------|----|-----|-----|-----|-------------|-----|------|-----|-----------------|------|------|-----|----------|-----|------|-----|-----|--|
| | | | | | | | | | | | | | | | | | | |
| WO 2008113606 | | | | | A1 20080925 | | | | 1 | WO 2 | 008- | | 20080321 | | | | | |
| | W: | ΑE, | AG, | AL, | AM, | AO, | AΤ, | ΑU, | ΑZ, | BA, | BB, | BG, | BH, | BR, | BW, | BY, | BZ, | |
| | | CA, | CH, | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DO, | DZ, | EC, | EE, | EG, | ES, | |
| | | FI, | GB, | GD, | GE, | GH, | GM, | GΤ, | HN, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | ΚE, | |
| | | KG, | KM, | KN, | KΡ, | KR, | KΖ, | LA, | LC, | LK, | LR, | LS, | LT, | LU, | LY, | MA, | MD, | |
| | | ME, | MG, | MK, | MN, | MW, | MX, | MY, | MZ, | NA, | NG, | NΙ, | NO, | NZ, | OM, | PG, | PH, | |
| | | PL, | PT, | RO, | RS, | RU, | SC, | SD, | SE, | SG, | SK, | SL, | SM, | SV, | SY, | ТJ, | TM, | |

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TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW
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             IE, IS, IT, LT, LU, LV, MC, MT, NL, NO, PL, PT, RO, SE, SI, SK,
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             TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW,
            AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
    US 20070269460
                         Α1
                               20071122
                                           US 2007-723638
                                                                  20070321
PRIORITY APPLN. INFO.:
                                           US 2007-723638
                                                              A2 20070321
                                           FR 2003-6772
                                                              A 20030605
                                           WO 2004-FR50214
                                                              W 20040604
                                           US 2005-559431
                                                              A2 20051205
AΒ
    The invention relates to the use of a therapeutically effective amt. of a
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peptide compn. comprising a polyprotein NS3/NS4 of the hepatitis C virus (HCV) as well as a polypeptide NS5b of the HCV, for the prepn. of a medicament for administration to a HCV-infected subject in the treatment of hepatitis C. The invention provides a compd. contq. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein NS3/NS4 and the polypeptide NS5b. Preclin. studies demonstrate that three sub-cutaneous injections of MVA vector-based NS34-NS5B construct at one week interval (weeks 1, 2 and 3) represent an optimized protocol to induce HCV specific IFNy producing T cells and cytotoxic T lymphocytes. A recall injection performed several months after the first series (either at month 4 or month 6) was able to enhance both CD4+ and CD8+ T cell responses.

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FUI 1ES

ACCESSION NUMBER: 2008:570045 CAPLUS

TITLE: A **vector**-based minigene vaccine approach results in

strong induction of T-cell responses specific of

hepatitis C virus

AUTHOR(S): Martin, Perrine; Simon, Benjamin; Lone, Yu-Chun;

Chatel, Laurence; Barry, Ronald; Inchauspe, Genevieve;

Fournillier, Anne

CORPORATE SOURCE: Infectious Diseases Department, TRANSGENE SA, Lyon,

69364, Fr.

SOURCE: Vaccine (2008), 26(20), 2471-2481

CODEN: VACCDE; ISSN: 0264-410X

PUBLISHER: Elsevier Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Summary: Multiepitope-based vaccines against hepatitis C virus (HCV) were designed in the form of three minigenes encompassing four domains of the NS3, NS4 and NS5B proteins that contain multiple class I/II restricted epitopes. The polyEp-WT minigene encodes all four domains in fusion, the polyEp-C minigene encodes the same fusion but optimized for mammalian translation and the polyEp-E3 minigene has an addnl. endoplasmic reticulum targeting sequence. Whereas the minigenes vectorised by DNA were poorly immunogenic, adenovirus vectorization induced strong and broader IFNy-ELISpot and CTL responses in HLA-A2 transgenic mice. In addn., polyEp-WT and polyEp-E3 responses were found cross-reactive in a recombinant Listeria-NS3-based surrogate challenge. This study illustrates the potency of vectorised minigenes in the field of HCV

vaccine development.

REFERENCE COUNT: 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FUI Text

ACCESSION NUMBER: 2007:1461700 CAPLUS

DOCUMENT NUMBER: 148:260241

TITLE: The Functional Evaluation of Dendritic Cell Vaccines

Based on Different Hepatitis C Virus Nonstructural

Genes

AUTHOR(S): Tian, Yuan; Zhang, Heng-Hui; Wei, Lai; Du, Shao-Cai;

Chen, Hong-Song; Fei, Ran; Liu, Feng

CORPORATE SOURCE: Hepatology Institute, Peking University People's

Hospital, Beijing, Peop. Rep. China

SOURCE: Viral Immunology (2007), 20(4), 553-561

CODEN: VIIMET; ISSN: 0882-8245

PUBLISHER: Mary Ann Liebert, Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

Hepatitis C virus (HCV) nonstructural (NS) genes are relatively conserved and play crit. roles in cellular immune responses against HCV. The aim of the study was to evaluate the immunogenicity of the different HCV NS genes through transduction of DCs and presentation to T cells. Monocyte-derived DCs from healthy donors were infected with the recombinant adenovirus (Ad) harboring HCV NS3 (AdNS3), NS4 (NS4A and NS4B; AdNS4), NS5 (NS5A and NS5B; AdNS5), NS3/NS4 (AdNS3/NS4), and NS4/NS5 (AdNS4/NS5) genes, and then used to stimulate autologous lymphocytes in vitro. Antigen-specific cellular immune responses were detected by interferon- γ (IFN- γ), interleukin 4 (IL-4), and Granzyme B (GrB) enzyme-linked immunospot assays (ELISPOT). expressing different HCV NS genes all induced pos. immune responses. Furthermore, DCs transfected with AdNS3/NS4 were superior to DCs infected with AdNS3 or AdNS4 in inducing HCV-specific immunity. The same results were obtained when the authors compared DCs infected with AdNS4/NS5 to AdNS4 or AdNS5. DCs transduced with NS3/NS4 or NS4/NS5

had similar ability to elicit specific immune responses to HCV. REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FUL

ACCESSION NUMBER: 2007:1334675 CAPLUS

DOCUMENT NUMBER: 148:9402

TITLE: Compositions comprising the hepatitis C virus (HCV)

polyprotein NS3/NS4 and protein NS5b,

recombinant expression and sequences thereof, and

vaccine uses

INVENTOR(S): Inchauspe, Genevieve; Fournillier, Anne

PATENT ASSIGNEE(S): Transgene S.A., Fr.

SOURCE: U.S. Pat. Appl. Publ., 75 pp., Cont.-in-part of U.S.

Ser. No. 559,431.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

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PATENT NO.
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    US 20070269460
                       A1 20071122 US 2007-723638
                                                                20070321
    FR 2855758
                       A1
                             20041210
                                         FR 2003-6772
                                                                20030605
    FR 2855758
                       В1
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                   A2
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    US 20060134065
                              20060622
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                        A1
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                                                           A 20030605
                                          FR 2003-6772
PRIORITY APPLN. INFO.:
                                          WO 2004-FR50214
                                                             W 20040604
                                          US 2005-559431
                                                           A2 20051205
                                          US 2007-723638
                                                            A2 20070321
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AB The invention provides a compd. contg. a polyprotein NS3/NS4 and a polypeptide NS5b of hepatitis C virus (HCV), which has an immunogenic and protective power superior to that obtained with a vaccine addnl. including the protein NS5a and/or other structural proteins of HCV. Said invention also relates to expression vectors, such as adenovirus and poxvirus vectors, encoding the polyprotein NS3/NS4 and the polypeptide NS5b. The inventive compd. can be used for a therapeutic application.

L5 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FULL
Text
ACCESSION NUMBER:

2007:1112932 CAPLUS

DOCUMENT NUMBER: 148:236736

TITLE: An acceler

An accelerated vaccine schedule with a poly-antigenic hepatitis C virus MVA-based candidate vaccine induces potent, long lasting and in vivo cross-reactive T cell

responses

AUTHOR(S): Fournillier, A.; Gerossier, E.; Evlashev, A.; Schmitt,

D.; Simon, B.; Chatel, L.; Martin, P.; Silvestre, N.;

Balloul, J. M.; Barry, R.; Inchauspe, G.

Site AFSSA, Transgene S.A., Lyon, 69364, Fr.

Vaccine (2007), 25(42), 7339-7353

CODEN: VACCDE; ISSN: 0264-410X

CORPORATE SOURCE: SOURCE:

PUBLISHER: Elsevier Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

AB We designed and evaluated in HLA-class I transgenic mouse models a hepatitis C virus (HCV) T cell-based MVA vectored vaccine expressing three viral antigens known to be targets of potent CD8+- and CD4+-mediated responses. An accelerated (3 wk-based) vaccination induced specific CD8+ T cells harboring two effector functions (cytolytic activity - both in vitro and in vivo - and prodn. of IFNγ) as well as specific CD4+ T cells recognizing all three vaccine antigens. Responses were long lasting (6 mo), boostable by a fourth MVA vaccination and in vivo cross-reactive as demonstrated in a surrogate Listeria-based challenge assay. This candidate vaccine has now moved into clin. trials.

REFERENCE COUNT: 61 THERE ARE 61 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

EU FEST

ACCESSION NUMBER: 2006:492228 CAPLUS

DOCUMENT NUMBER: 144:487147

TITLE: Yeast-based therapeutic vaccine vehicle for chronic

hepatitis c infection

INVENTOR(S): Duke, Richard C.; Franzusoff, Alex; Haller, Aurelia;

King, Thomas H.

PATENT ASSIGNEE(S): Globeimmune, Inc., USA

SOURCE: U.S. Pat. Appl. Publ., 47 pp., Cont.-in-part of U.S.

Ser. No. 738,646.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE | | |
|------------------------|------|----------|-----------------|----|----------|--|--|
| | | | | | | | |
| US 20060110755 | A1 | 20060525 | US 2005-254252 | | 20051018 | | |
| <u>US 7439042</u> | В2 | 20081021 | | | | | |
| <u>US 20040156858</u> | A1 | 20040812 | US 2003-738646 | | 20031216 | | |
| US 7465454 | B2 | 20081216 | | | | | |
| <u>US 20080069833</u> | A1 | 20080320 | US 2007-768144 | | 20070625 | | |
| PRIORITY APPLN. INFO.: | | | US 2002-434163P | Р | 20021216 | | |
| | | | US 2003-738646 | A2 | 20031216 | | |
| | | | US 2004-620158P | Р | 20041018 | | |

OTHER SOURCE(S): MARPAT 144:487147

AB The present invention relates to compns., including vaccines, and methods for vaccinating an animal against hepatitis C virus (HCV) and for treating or preventing hepatitis C viral infection in an animal. The invention includes a variety of novel HCV fusion proteins that can be used directly as a vaccine or in conjunction with a yeast-based vaccine vehicle to elicit an immune response against HCV in an animal. The invention also includes the use of the HCV fusion gene and protein described herein in any diagnostic or therapeutic protocol for the detection and/or treatment or prevention of HCV infection.

L5 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

EJII Text

ACCESSION NUMBER: 2006:333454 CAPLUS

DOCUMENT NUMBER: 144:357638

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TITLE:
                          Application of a transgenic mouse model of hepatitis c
                          virus (HCV) infection and identification of
                          antiviral agent for HCV therapeutics
INVENTOR(S):
                          Sallberg, Matti; Frelin, Lars
                          Tripep AB, Swed.
PATENT ASSIGNEE(S):
SOURCE:
                          PCT Int. Appl., 165 pp.
                          CODEN: PIXXD2
DOCUMENT TYPE:
                          Patent
LANGUAGE:
                          English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:
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                                 DATE APPLICATION NO. DATE
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     WO 2006021896
                          A2
                                20060302
                                             WO 2005-IB3736
                                                                       20050826
     WO 2006021896
                          A3 20060817
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                          A2 20070509
                                             EP 2005-810181
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                      A2
     WO 2006109196
                               20061019
                                             WO 2006-IB1668
                          AЗ
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     WO 2006109196
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                      A1 20081127
                                              US 2008-660878
     US 20080295185
                                                                       20080506

      US
      2004-605030P
      P
      20040827

      US
      2005-649975P
      P
      20050204

      WO
      2005-IB3736
      W
      20050826

      US
      2005-740362P
      P
      20051128

PRIORITY APPLN. INFO.:
AΒ
     Disclosed herein is the discovery of novel NS3/4A compns. with enhanced
     expression abilities. Embodiments of the invention include codon
     optimized NS3/4A compns. and compns. with the Semliki forest virus
     replicon. Addnl. embodiments include transgenic organisms contg. these
     NS3/4A compns., methods of using these transgenic mice to screen and
     refine drugs, and the drugs refined by these methods. Addnl. embodiments
     include protease activity dependent mols. that can indicate the presence
     or absence of a protease inhibitor.
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http://stnweb.cas.org/cgi-bin/sdcgi?SID=858997-1982474537-200&APP=stnweb&

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

3

REFERENCE COUNT:

L5 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

Fell Text

ACCESSION NUMBER: 2005:303181 CAPLUS

DOCUMENT NUMBER: 142:372468

TITLE: HCV fusion proteins with modified NS3 domains and

uses thereof as immunogens

INVENTOR(S): Houghton, Michael

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S.

Ser. No. 721,479.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|----|----------|
| | | | | | |
| US 20050074465 | A1 | 20050407 | US 2003-612884 | | 20030702 |
| <u>US 6986892</u> | B1 | 20060117 | US 2000-721479 | | 20001122 |
| US 20060057164 | A1 | 20060316 | US 2005-195009 | | 20050802 |
| <u>US 7449566</u> | B2 | 20081111 | | | |
| JP 2006265267 | A | 20061005 | JP 2006-174595 | | 20060623 |
| PRIORITY APPLN. INFO.: | | | US 1999-167502P | P | 19991124 |
| | | | US 2000-721479 | Α2 | 20001122 |
| | | | US 2002-393694P | P | 20020702 |
| | | | US 2002-394510P | P | 20020708 |
| | | | JP 2004-519849 | АЗ | 20030702 |

The disclosed invention provides hepatitis C virus (HCV) fusion proteins that include a mutated NS3 protease domain, fused to at least one other HCV epitope derived from another region of the HCV polyprotein. The fusions can be used in stimulation of a cellular immune response to HCV, such as activating hepatitis C virus (HCV)-specific T cells, including CD4+ and CD8+ T cells. The method can be used in model systems to develop HCV-specific immunogenic compns., as well as to immunize a mammal against HCV. In expts. with Rhesus macaques, the immunization with plasmid DNA encoding an NS3 (modified) NS4NS5aCore fusion protein led to activation of HCV-specific CD8-pos. T cells expressing interferon γ and proliferation of HCV-specific CD4-pos. T cells. Also presented is the use of alphavirus replicon particles.

L5 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FUIL TEXT

ACCESSION NUMBER: 2004:905910 CAPLUS

DOCUMENT NUMBER: 141:378844

TITLE: Inducing a T cell response with recombinant

antigen-expressing pestivirus replicons or recombinant pestivirus replicon-transfected dendritic cells, and

therapeutic uses

INVENTOR(S): Rehermann, Barbara; Racanelli, Vito; Behrens,

Sven-Erik; Tautz, Norbert

PATENT ASSIGNEE(S): The Government of the United States of America as

Represented by the Secretary of Health and Human Services, USA; Justus-Liebig-Universitaet Giessen

SOURCE: PCT Int. Appl., 143 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| | PATENT NO. | | | | | KIND DATE | | | | APPL | | DATE | | | | | | |
|------|--------------------------------|-----|-----|-----|----------|-----------|----------|------------|-----|-----------------|------|------|------|-----|----------|-----|------|-----|
| | WO 2004092386 WO 2004092386 | | | | A2 A3 | | 20041028 | | 1 | WO 2004-US11018 | | | | | 20040410 | | | |
| | WO_ | W: | AE, | AG, | | AM, | AT, | AU, DE, | AZ, | | | | | | | | • | |
| | | | GE, | GH, | GM, | HR, | HU, | ID, LV, | IL, | IN, | IS, | JP, | KE, | KG, | KP, | KR, | KZ, | LC, |
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AB The present disclosure relates to compds. and methods of generating T cell-mediated immunity, particularly T cell-mediated immunity to Hepatitis C Virus (HCV), Respiratory Syncytial Virus (RSV), Human Immunodeficiency Virus (HIV), Mycobacterium tuberculosis, Plasmodium falciparum, and tumors. The method includes (a) administering to the subject an amt. of an antigen presenting cell (such as dendritic cell) sufficient to induce the response in the subject, wherein the antigen presenting cell expresses the recombinant antigen from a pestivirus replicon or (b) directly administering the recombinant antigen expressing replicon in form of RNA or DNA.

REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FUI TEXT

ACCESSION NUMBER: 2002:908392 CAPLUS

DOCUMENT NUMBER: 138:13314

TITLE: Comparative vaccine studies in HLA-A2.1-transgenic

mice reveal a clustered organization of epitopes presented in hepatitis C virus natural infection

AUTHOR(S): Himoudi, Nourredine; Abraham, Jean-Daniel;

Fournillier, Anne; Lone, Yu Chun; Joubert, Aurelie; Op De Beeck, Anne; Freida, Delphinc; Lemonnier, Francois;

Kieny, Marie Paule; Inchauspe, Genevieve

CORPORATE SOURCE: Unite Mixte CNRS-BioMerieux, UMR 2142, Ecole Normale

Superieure, Lyon, 69364, Fr.

SOURCE: Journal of Virology (2002), 76(24), 12735-12746

CODEN: JOVIAM; ISSN: 0022-538X

PUBLISHER: American Society for Microbiology

DOCUMENT TYPE: Journal LANGUAGE: English

AB A polyepitopic CD8+-T-cell response is thought to be crit. for control of hepatitis C virus (HCV) infection. Using transgenic mice, we analyzed the immunogenicity and dominance of most known HLA-A2.1 epitopes presented during infection by using vaccines that carry the potential to enter clin. trials: peptides, DNA, and recombinant adenoviruses. The vaccines capacity to induce specific cytotoxic T lymphocytes and interferon gamma-producing cells revealed that immunogenic epitopes are clustered in

specific antigens. For two key antigens, flanking regions were shown to greatly enhance the scope of epitope recognition, whereas a DNA-adenovirus prime-boost vaccination strategy augmented epitope immunogenicity, even that of subdominant ones. The present study reveals a clustered organization of HCV immunogenic HLA.A2.1 epitopes and strategies to modulate their dominance.

REFERENCE COUNT:

51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 11 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

Text

2002:716438 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 137:227663

TITLE: Hepatitis C virus (HCV) cDNA-based hepatocyte cell

culture system for synthesis of infectious HCV, and

uses for antiviral screening

INVENTOR(S): Dasgupta, Asim; Koka, Prasad S.

PATENT ASSIGNEE(S): The Regents of the University of California, USA

SOURCE: PCT Int. Appl., 42 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PAT | PATENT NO. | | | | | D | DATE | | APPLICATION NO. | | | | | DATE | | | | |
|--------------------|------------|-------|-----|-----|-----------------|----------|------|----------------|-----------------------|------|------|------|----------|----------|-----|-----|-----|--|
| WO | 2002 | 0727 | 76 | | A2 | | 2002 | 0919 | WO 2002-US7516 | | | | | 20020311 | | | | |
| WO | 2002 | 0727 | 76 | | A3 2004020 | | | 0205 | | | | | | | | | | |
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| | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | ES, | FI, | GB, | GD, | GE, | GH, | |
| | | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, | KG, | KΡ, | KR, | KΖ, | LC, | LK, | LR, | |
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| | | GN, | GQ, | GW, | \mathtt{ML} , | MR, | NΕ, | SN, | TD, | TG | | | | | | | | |
| CA | 2440 | 433 | | | A1 20020919 | | | | | CA 2 | 002- | 2440 | 433 | | | | | |
| | | | | | | | | | <u>AU 2002-254190</u> | | | | | | | | | |
| US | 2002 | 0197: | | | | | | | <u>US 2002-96039</u> | | | | | 20020311 | | | | |
| | 7183 | | | | | | 2007 | | | | | | | | | | | |
| EΡ | 1421 | | | | | | 2004 | | | | | | | | | | | |
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| | | • | • | • | • | • | RO, | • | • | • | | | | | | | | |
| | | | | | 20041216 | | | JP 2002-571832 | | | | | 20020311 | | | | | |
| <u>CN 1592794</u> | | | | А | | 20050309 | | | CN 2002-806237 | | | | | | | | | |
| RITY APPLN. INFO.: | | | .: | | | | | | US 2 | | | | | | | | | |
| mb | | | | | | | | | | WO 2 | | | | | | | 311 | |

The present invention presents a method of synthesizing infectious AΒ hepatitis C virus (HCV) by transfecting hepatocyte cells with a gene encoding HCV and then exposing uninfected cells to the HCV to form addnl. $\mbox{HCV}.$ The invention relates to a \mbox{HCV} cDNA-based culture system capable of synthesis of infectious HCV in cell culture and cell-to-cell spread of the virus. The expression of T7 RNA polymerase in the cytoplasm was used to transcribe the HCV cDNA under the T7 promoter to generate high quantities of HCV RNA. The viral RNA proved to be translated to produce viral structural (core, El, E2 and p7) and nonstructural (NS2,

NS3, NS4A and B, NS5A and B) proteins. Viral RNA replication directed by the RNA-dependent RNA polymerase (NS5B) would then occur. Progeny virions were made and secreted into the tissue culture media, and infection of neighboring cells resulting in cell-to-cell spread of virus was demonstrated. The invention also relates to a method of measuring the level of HCV infection in a hepatocyte cell. A method for identifying a modulator of HCV activity is also presented, and a method for modulating HCV activity. The invention provides a reliable system for both genetic anal. of the viral genome and for the development of novel antiviral strategies.

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FUI Text ACCESSION NUMBER:

ACCESSION NUMBER: 1999:113845 CAPLUS

DOCUMENT NUMBER: 130:163166

TITLE: Test **vectors** containing hepatitis C or human

cytomegalovirus nucleic acid and indicator gene and methods for determining antiviral susceptibility and

resistance and for antiviral screening

INVENTOR(S): Capon, Daniel J.; Whitcomb, Jeannette M.; Parkin, Neil

Τ.

PATENT ASSIGNEE(S): Virologic, Inc., USA

SOURCE: PCT Int. Appl., 128 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| | PATENT NO. | | | | | KIND DATE | | | | | APPL | ICAT | ION 1 | DATE | | | | |
|------|------------------------|------|------|-----------|-------------|-----------|-----------------|------|----------------|------|----------|-------------|-------|------------|-----|-------|------|-----|
| | WO 9906597 | | | | | | WO 1998-US15967 | | | | | 19980730 | | | | | | |
| | | W: | AL, | AM, | AT, | ΑU, | AZ, | BA, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CU, | CZ, | DE, |
| | | | DK, | EE, | ES, | FI, | GB, | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IS, | JP, | KΕ, | KG, |
| | | | KΡ, | KR, | KΖ, | LC, | LK, | LR, | LS, | LT, | LU, | LV, | MD, | MG, | MK, | MN, | MW, | MX, |
| | | | NO, | NΖ, | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, |
| | | | UA, | UG, | UZ, | VN, | YU, | ZW | | | | | | | | | | |
| | | RW: | GH, | GM, | ΚE, | LS, | MW, | SD, | SZ, | UG, | ZW, | ΑT, | BE, | CH, | CY, | DE, | DK, | ES, |
| | | | FI, | FR, | GB, | GR, | IE, | ΙT, | LU, | MC, | NL, | PT, | SE, | BF, | ВJ, | CF, | CG, | CI, |
| | | | CM, | GΑ, | GN, | GW, | \mathtt{ML} , | MR, | NE, | SN, | TD, | TG | | | | | | |
| | CA | 2298 | 102 | | | A1 | | 1999 | 0211 | 1 | CA 1 | 998- | 2298: | 102 | | 1 | 9980 | 730 |
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| | EP 1012334 | | | A1 | A1 20000628 | | | | EP 1 | 9407 | 19980730 | | | | | | | |
| | | R: | ΑT, | BE, | CH, | DE, | DK, | ES, | FR, | GB, | GR, | ΙT, | LI, | LU, | NL, | SE, | MC, | PT, |
| | | | IE, | FΙ | | | | | | | | | | | | | | |
| | JP | 2001 | 5120 | <u>36</u> | | T | | 2001 | 0821 | | JP 2 | 000- | 5053 | 36 | | 1 | 9980 | 730 |
| PRIO | PRIORITY APPLN. INFO.: | | | | | | | | US 1997-903507 | | | | | A 19970730 | | | | |
| | | | | | | | | | 1 | WO 1 | 998-1 | <u>US15</u> | 967 | Ţ | W 1 | 9980. | 730 | |

This invention provides a method for detg. susceptibility for an **HCV** or HCMV anti-viral drug comprising: (a) introducing a resistance test **vector** comprising a patient-derived segment and an indicator gene into a host cell; (b) culturing the host cell from (a); (c) measuring expression of the indicator gene in a target host cell, and (d) comparing the expression of the indicator gene from (c) with the expression of the indicator gene measured when steps (a-c) are carried out in the absence of the anti-viral drug, wherein a test concn. of the anti-viral drug is present at steps (a-c); at steps (b-c); or at step (c). This invention

also provides a method for detg. HCV or HCMV anti-viral drug resistance in a patient comprising: (a) detg. anti-viral drug susceptibility in the patient at a first time using the susceptibility test described above, wherein the patient-derived segment is obtained from the patient at about said time; (b) detg. anti-viral drug susceptibility of the same patient at a later time; and (c) comparing the anti-viral drug susceptibilities detd. in step (a) and (b), wherein a decrease in anti-viral drug susceptibility at the later time compared to the first time indicates development or progression of anti-viral drug resistance in the patient. This invention also provides a method for evaluating the biol. effectiveness of a candidate HCV or HCMV anti-viral drug compd. Compns. including resistance test vectors comprising a patient-derived segment comprising an HCV or HCMV gene and an indicator gene and host cells transformed with the resistance test vectors are provided.

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

EGI TEXE

ACCESSION NUMBER: 1998:251284 CAPLUS

DOCUMENT NUMBER: 128:292153

ORIGINAL REFERENCE NO.: 128:57803a,57806a

TITLE: Protease regulator screening assay using a recombinant

polypeptide comprising anchor, protease recognition,

and signal regions

INVENTOR(S): Chien, David Y.; Selby, Mark J.

PATENT ASSIGNEE(S): Chiron Corporation, USA SOURCE: PCT Int. Appl., 41 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| | PATENT NO. | | | | | KIND DATE | | | | APPLICATION NO. | | | | | | DATE | | | |
|------|------------------------|-------|------|-----|-----|-------------|------------|------|-----------------|----------------------|------|-------|-----|------------|-------------|------|------|-----|--|
| | WO 9816657 | | | | A1 | A1 19980423 | | | WO 1997-US18632 | | | | | 19971017 | | | | | |
| | W: AL, AM, A | | | ΑT, | AU, | AZ, | BA, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CU, | CZ, | DE, | | |
| | | | DK, | EE, | ES, | FΙ, | GB, | GE, | GH, | HU, | ID, | IL, | IS, | JP, | KE, | KG, | KΡ, | KR, | |
| | | | KΖ, | LC, | LK, | LR, | LS, | LT, | LU, | LV, | MD, | MG, | MK, | MN, | MW, | MX, | NO, | NZ, | |
| | | | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, | UA, | UG, | |
| | | | UZ, | VN, | YU, | ZW | | | | | | | | | | | | | |
| | | RW: | GH, | KE, | LS, | MW, | SD, | SZ, | UG, | ZW, | AΤ, | BE, | CH, | DE, | DK, | ES, | FI, | FR, | |
| | | | GB, | GR, | ΙE, | ΙT, | LU, | MC, | ΝL, | PT, | SE, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | |
| | | | GN, | ML, | MR, | NE, | SN, | TD, | TG | | | | | | | | | | |
| | ΑU | 9749 | 043 | | | Α | | 1998 | 0511 | <u>AU 1997-49043</u> | | | | | | 1 | 9971 | 017 | |
| | US | 6436 | 666 | | | В1 | | 2002 | 0820 | US 1997-997055 | | | | | 19971017 | | | | |
| | US | 2003 | 0113 | 825 | | A1 | 1 20030619 | | | US 2002-225390 | | | | | 20020820 | | | | |
| | US | 69241 | 122 | | | В2 | | 2005 | 0802 | | | | | | | | | | |
| | | | | | A1 | | 2006 | 1228 | US 2005-193615 | | | | | | 20050801 | | | | |
| | US | 7439 | 040 | | | В2 | | 2008 | 1021 | | | | | | | | | | |
| PRIO | PRIORITY APPLN. INFO.: | | | . : | | | | | | US 1 | 996- | 2881 | 7 P | | P 1 | 9961 | 017 | | |
| | | | | | | | | | | US 1 | 997- | 9970. | 55 | | A1 1 | 9971 | 017 | | |
| | | | | | | | | | WO 1997-US18632 | | | | | W 19971017 | | | | | |
| | | | | | | | | | | US 2002-225390 | | | | | A3 20020820 | | | | |
| | | | | | | | _ | | | | | | | | | | _ | | |

AB A polypeptide contg. an anchor region, a protease recognition site, and a detectable signal region can be produced recombinantly and directly attached to a solid support. The polypeptide is useful for screening protease regulators, esp. protease inhibitors. Thus, a recombinant

protein is produced in which the anchor region is protein A which specifically binds to an antibody, the protease recognition site is that for hepatitis C virus NS3 protease such as that for NS4A/NS4B or HS4B/NS5A cleavage, and the signal region comprises the epitope FLAG sequence. A fragment encoding HCV NS5 peptide protease target site is inserted in frame into the polylinker region of pEZZ18 so that it is connected at the C-terminal region of protein A. The NS5 peptide protease target site includes the NS5A and NS5B cleavage site, i.e., amino acids 2420 and 2421, 7 amino acids at the N-terminal side of the cleavage site, and 8 amino acids at the C-terminal side of the cleavage site. Another sequence fragment encoding the FLAG tag is inserted in frame at the C-terminal end of the NS5 protease target site. The sequence fragment encodes three FLAG tags alternately spaced with two 4-glycine spacers. The assay is readily adapted to an automated format and is suitable for large scale drug screens, as demonstrated by screening for potentially therapeutically useful inhibitors of the HCV protease.

REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

FUI TEXE

PUBLISHER:

ACCESSION NUMBER: 1997:228414 CAPLUS

DOCUMENT NUMBER: 126:247257

ORIGINAL REFERENCE NO.: 126:47707a,47710a

TITLE: Hepatitis C virus (HCV) RNA polymerase assay using

cloned HCV non-structural proteins

AUTHOR(S): Bartholomeusz, Angeline I.; Guo, Ke-Jian; Edwards,

Patrick C.; Locarnini, Stephen A.

CORPORATE SOURCE: Victorian Infectious Diseases Reference Laboratory,

Victoria, 3078, Australia

SOURCE: Antiviral Therapy (1996), 1(Suppl. 4, Therapies for

Viral Hepatitis), 18-24

CODEN: ANTHFA; ISSN: 1359-6535 International Medical Press

Investigations into the RNA replication of hepatitis C virus (HCV) have

DOCUMENT TYPE: Journal LANGUAGE: English

been hampered by the lack of a cell-culture system. The objective of this study was to develop an in vitro system to study **HCV** polymerase activity and RNA replication. We are currently developing two **HCV** RNA replication assays. The first reconstitutes the various components required for RNA synthesis: cloned viral non-structural proteins as the source of the viral polymerase and helicase, exts. from uninfected Vero (African green monkey kidney) or HepG2 (human hepatoma) cells as the source of host factors and an RNA template (either **HCV** RNA transcripts or RNA from the pestivirus bovine viral diarrhea virus). The second assay uses **HCV**-infected liver cell exts. and thus contains authentic replication complexes consisting of viral and host proteins and RNA templates. In both assays, synthesis of viral RNA is detected by the incorporation of the radiolabel [α -32P]GTP. In the assay using cloned viral protein, the genes encoding NS2, **NS3**, **NS4**, NS5A and **NS5B** from pBRTM/**HCV** 1-3011 were cloned into the transcription **vector**

pT7T3. The transcribed RNA was translated with rabbit reticulocytes in the presence of canine pancreatic membranes. Radiolabeled RNA was detected only in polymerase assays that contained the translated proteins and all other components. In assays using infected liver cell exts., radiolabel was incorporated into RNA products that were not present in control assays using uninfected liver cell exts. Both assays will be useful in the elucidation of processes involved in **HCV** RNA replication

and in the development of antiviral agents.

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